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ATTORNEY'S DOCKET NUMBER

TRANSMITTAL LETTER TO THE UNITED STATES B-0497 PUS TON 10 Of known, see 37 CFR 1.5) **DESIGNATED/ELECTED OFFICE (DO/EO/US)** CONCERNING A SUBMISSION UNDER 35 U.S.C. 371 INTERNATIONAL APPLICATIOAN NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED PCT/EP2003/012514 November 10, 2003 November 13, 2002 TITLE OF INVENTION METHOD FOR THE SELECTIVE AND QUANTITATIVE FUNCTIONALIZATION OF IMMUNOGLOBULIN FAB FRAGMENTS. CONJUGATE COMPOUNDS OBTAINED WITH THE SAME AND COMPOSITION THEREOF APPLICANT(S) FOR DO/ES DE HAEN, Christoph and MAISANO, Federico Applicant herewith submits to the united States Designated/Elected Office (DO/EO/S) the following items and other information: This is a FIRST submission of items concerning a submission under 35 U.S.C. 371. 2. This is a SECOND or SUBSEQUENT submission of items concerning a submission under 35 U.S.C. 371. 3. This is an express request to begin national examination procedures (335 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below The US has been elected by the expiration of 19 months from the priority date (Article 31). 4 5. A copy of the International Application as filed (35 US.C. 371(c)(2)) is attached hereto (required only if not communicated by the International Bureau). X b. has been communicated by the International Bureau. is not required, as the application was filed in the United States Receiving Office (RO/US). C. An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). 6. a. is attached hereto. has been previously submitted under 35 U.S.C. 154(d)(4). b. Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3)). 7. are attached hereto (required only if not communicated by the International Bureau). have been communicated by the International Bureau. C. have not been made; however, the time limit for making such amendments has NOT expired. have not been made and will not be made. An English language translation of the annexes of the International Preliminary Examination report under PCT Article 36 (35 U.S.C. 371(c)(3)). g An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)). An English translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)). 10. Items 11 to 20 below concern document(s) or information included: X An Information Disclosure Statement under 37 CFR 1.97 and 198. 11. X 12. An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. 13. X A preliminary amendment. 14. X An Application Data Sheet under 37 CFR 1.76. 15. A substitute specification. 16. ΙX A power of attorney and/or change of address letter. A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821-1.825. 17. 18. A second copy of the published International Application under 35 US.C. 154(d)(4). 19. A second copy of the English language translation of the International Application under 35 U.S.C. 154(d)(4). 20 X Other items or information: Copy of the PCT Publication; International Examination Report; PTO-1449 & 9 cited references and Statement Under 37 CFR 3.73(b)

This collection of information is required by 37 CFR 1.414 and 1.491-1.492. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.11 and 1.14. This collection is estimated to take 15 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require completing this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NO SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Mail Stop PCT, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

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INTERNATIONAL APPLICATION NO. U.S. APPLICATION NO. (if known, see 37 CFR 1.5) PCT/EP2003/012514 B-0497 PUS **CALCULATIONS** The following fees have been submitted PTO USE ONLY \$300 21. X Basic national fee.....\$300 Examination fee If International preliminary examination report prepared by USPTO and all claims satisfy provisions of \$200 23. X Search fee Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the USPTO as an International Searching Authority.....\$100 \$400 All other situations.....\$500 \$900 TOTAL OF 21, 22 and 23 = Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets of paper or fraction thereof. **Total Sheets** Extra Sheets Number of each additional 50 or fraction RATE thereof (round up to a whole number) 35 - 100 = 0 /50 =x \$250 \$000 Surcharge of \$130.00 for furnishing the oath or declaration later than 30 months from the earliest \$ claimed priority date (37 CFR 1.492(h)). NUMBER FILED **CLAIMS** NUMBER EXTRA **RATE** \$ 22 2 -20 =x \$50 \$100 Total claims 0 Independent claims x \$200 \$000 MULTIPLE DEPENDENT CLAIM(S) (if applicable) \$360 \$362 \$462 TOTAL OF ABOVE CALCULATIONS = Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2. \$ 1362 Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest \$ claimed priority date (37 CFR 1.492(i)). **TOTAL NATIONAL FEE =** \$13<u>62</u> Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied \$ 40 by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property TOTAL FEES ENCLOSED = \$ 1402 Amount to be \$ refunded: Amount to be \$1,402.00 charged: A check in the amount of \$ ___ _____ to cover the above fees is enclosed. Please charge my Deposit Account No. 50-2168 in the amount of $\frac{1,402.00}{}$ to cover the above fees. A duplicate copy of this sheet is enclosed. The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit _. A duplicate copy of this sheet is enclosed. Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038. NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the International Application to pending status. SEND ALL CORRESPONDENCE TO: ATTN: M. Caragh Noone, Esq. Bracco Research USA Inc. Princeton, NJ 08540